IN THE UNITED STATES PATENT AND TRADEMARK OFFICE



In re Patent Application of

Elmore et al

Atty. Ref.: 1498-133

Serial No. 08/981,087

Group: 1647

Filed: May 27, 1998

Examiner: Turner

For: TYPE F BOTULINUM TOXIN AND USE THEREOF

January 28, 2003

Assistant Commissioner for Patents Washington, DC 20231

Sir:

STATEMENT

The attached paper and computer readable copies of the Sequence Listing are the same. No new matter has been added.

Respectfully submitted,

NIXON & VANDERHYE P.C.

Ву

B. J. Sadoff

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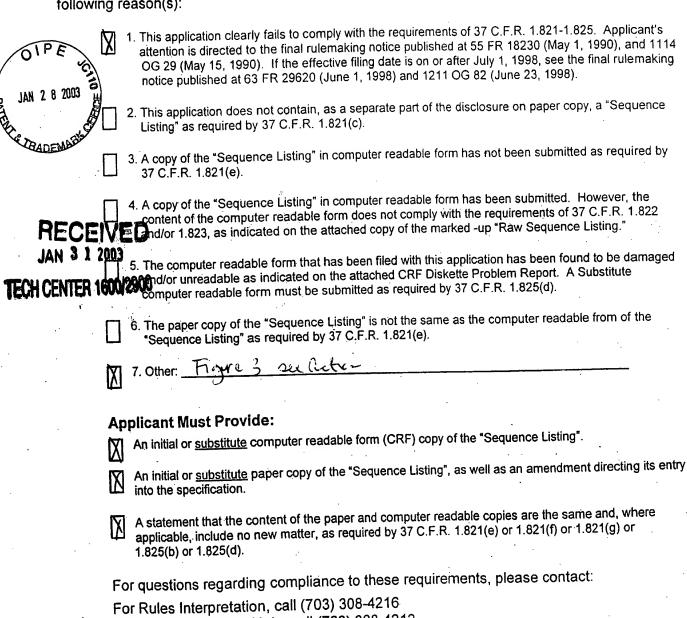
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Application No.:	
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NOTICE TO COMPLY WITH REQUIREMENTS FOR PATENT APPLICATIONS CONTAINING NUCLEOTIDE SEQUENCE AND/OR AMINO ACID SEQUENCE DISCLOSURES

Applicant must file the items indicated below within the time period set the Office action to whic' the Notice is attached to avoid abandonment under 35 U.S.C. § 133 (extensions of time may bootained under the provisions of 37 CFR 1.136(a)).

The nucleotide and/or amino acid sequence disclosure contained in this application does not comply with the requirements for such a disclosure as set forth in 37 C.F.R. 1.821 - 1.825 for the following reason(s):



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SEQUENCE LISTING

(1) GENERAL INFORMATION:

- (ii) TITLE OF INVENTION: TYPE F BOTULINUM TOXIN AND USE THEREOF
- (iii) NUMBER OF SEQUENCES: 8
- (iv) CORRESPONDENCE ADDRESS:
 - (A) ADDRESSEE: NIXON & VANDERHYE P.C.
 - (B) STREET: 1100 North Glebe Rd. 8th floor
 - (C) CITY: Arlington
 - (D) STATE: VA
 - (E) COUNTRY: USA
 - (F) ZIP: 22201-4741
- (v) COMPUTER READABLE FORM:
 - (A) MEDIUM TYPE: Floppy disk
 - (B) COMPUTER: IBM PC compatible
 - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
 - (D) SOFTWARE: PatentIn Release #1.0, Version #1.30
- (vi) CURRENT APPLICATION DATA:
 - (A) APPLICATION NUMBER: US 08/981,087
 - (B) FILING DATE: 27-MAY-1998
 - (C) CLASSIFICATION:
- (vii) PRIOR APPLICATION DATA:
 - (A) APPLICATION NUMBER: PCT/GB96/01409
 - (B) FILING DATE: 12-JUN-1996
- (vii) PRIOR APPLICATION DATA:
 - (A) APPLICATION NUMBER: GB 9511909.5
 - (B) FILING DATE: 12-JUN-1995
- (viii) ATTORNEY/AGENT INFORMATION:
 - (A) NAME: Crawford, Arthur R.
 - (B) REGISTRATION NUMBER: 25,327
 - (C) REFERENCE/DOCKET NUMBER: 124-688
 - (ix) TELECOMMUNICATION INFORMATION:
 - (A) TELEPHONE: 703-816-4000
 - (B) TELEFAX: 703-816-4100
- (2) INFORMATION FOR SEQ ID NO:1:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 431 amino acids
 - (B) TYPE: amino acid

- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:1:

Ser Tyr Thr Asn Asp Lys Ile Leu Ile Leu Tyr Phe Asn Lys Leu Tyr 1 5 10 15

Lys Lys Ile Lys Asp Asn Ser Ile Leu Asp Met Arg Tyr Glu Asn Asn 20 25 30

Lys Phe Ile Asp Ile Ser Gly Tyr Gly Ser Asn Ile Ser Ile Asn Gly 35 40 45

Asp Val Tyr Ile Tyr Ser Thr Asn Arg Asn Gln Phe Gly Ile Tyr Ser 50 55 60

Ser Lys Pro Ser Glu Val Asn Ile Ala Gln Asn Asn Asp Ile Ile Tyr 65 70 75 80

Asn Gly Arg Tyr Gln Asn Phe Ser Ile Ser Phe Trp Val Arg Ile Pro 85 90 95

Lys Tyr Phe Asn Lys Val Asn Leu Asn Asn Glu Tyr Thr Ile Ile Asp 100 105 110

Cys Ile Arg Asn Asn Asn Ser Gly Trp Lys Ile Ser Leu Asn Tyr Asn 115 120 125

Lys Ile Ile Trp Thr Leu Gln Asp Thr Ala Gly Asn Asn Gln Lys Leu 130 135 140

Val Phe Asn Tyr Thr Gln Met Ile Ser Ile Ser Asp Tyr Ile Asn Lys 145 150 155 160

Trp Ile Phe Val Thr Ile Thr Asn Asn Arg Leu Gly Asn Ser Arg Ile 165 170 175

Tyr Ile Asn Gly Asn Leu Ile Asp Glu Lys Ser Ile Ser Asn Leu Gly
180 185 190

Asp Ile His Val Ser Asp Asn Ile Leu Phe Lys Ile Val Gly Cys Asn 195 200 205

Asp Thr Arg Tyr Val Gly Ile Arg Tyr Phe Lys Val Phe Asp Thr Glu 210 215 220

Leu Gly Lys Thr Glu Ile Glu Thr Leu Tyr Ser Asp Glu Pro Asp Pro 225 230 235 240

Ser Ile Leu Lys Asp Phe Trp Gly Asn Tyr Leu Leu Tyr Asn Lys Arg 245 250 255

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Tyr Tyr Leu Leu Asn Leu Leu Arg Thr Asp Lys Ser Ile Thr Gln Asn 260 265 270

Ser Asn Phe Leu Asn Ile Asn Gln Gln Arg Gly Val Tyr Gln Lys Pro 275 280 285

Asn Ile Phe Ser Asn Thr Arg Leu Tyr Thr Gly Val Glu Val Ile Ile 290 295 300

Arg Lys Asn Gly Ser Thr Asp Ile Ser Asn Thr Asp Asn Phe Val Arg 305 310 315 320

Lys Asn Asp Leu Ala Tyr Ile Asn Val Val Asp Arg Asp Val Glu Tyr 325 330 335

Arg Leu Tyr Ala Asp Ile Ser Ile Ala Lys Pro Glu Lys Ile Ile Lys 340 345 350

Leu Ile Arg Thr Ser Asn Ser Asn Ser Leu Gly Gln Ile Ile Val 355 360 365

Met Asp Ser Ile Gly Asn Asn Cys Thr Met Asn Phe Gln Asn Asn Asn 370 $$ 375 $$ 380

Gly Gly Asn Ile Gly Leu Leu Gly Phe His Ser Asn Asn Leu Val Ala 385 390 395 400

Ser Ser Trp Tyr Tyr Asn Asn Ile Arg Lys Asn Thr Ser Ser Asn Gly \$405\$ \$410\$

Cys Phe Trp Ser Phe Ile Ser Lys Glu His Gly Trp Gln Glu Asn 420 425 430

(2) INFORMATION FOR SEQ ID NO:2:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 144 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:2:

Ser Tyr Thr Asn Asp Lys Ile Leu Ile Leu Tyr Phe Asn Lys Leu Tyr 1 5 10 15

Lys Lys Ile Lys Asp Asn Ser Ile Leu Asp Met Arg Tyr Glu Asn Asn 20 25

Lys Phe Ile Asp Ile Ser Gly Tyr Gly Ser Asn Ile Ser Ile Asn Gly 35 40 45

H2

Asp Val Tyr Ile Tyr Ser Thr Asn Arg Asn Gln Phe Gly Ile Tyr Ser 50 55 60

Ser Lys Pro Ser Glu Val Asn Ile Ala Gln Asn Asn Asp Ile Ile Tyr 65 70 75 80

Asn Gly Arg Tyr Gln Asn Phe Ser Ile Ser Phe Trp Val Arg Ile Pro 85 90 95

Lys Tyr Phe Asn Lys Val Asn Leu Asn Asn Glu Tyr Thr Ile Ile Asp 100 105 110

Cys Ile Arg Asn Asn Asn Ser Gly Trp Lys Ile Ser Leu Asn Tyr Asn 115 120 125

Lys Ile Ile Trp Thr Leu Gln Asp Thr Ala Gly Asn Asn Gln Lys Leu 130 135 140

(2) INFORMATION FOR SEQ ID NO:3:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 144 amino acids

(B) TYPE: amino acid

(C) STRANDEDNESS:

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO:3:

Val Phe Asn Tyr Thr Gln Met Ile Ser Ile Ser Asp Tyr Ile Asn Lys

1 10 15

Trp Ile Phe Val Thr Ile Thr Asn Asn Arg Leu Gly Asn Ser Arg Ile 20 25 30

Tyr Ile Asn Gly Asn Leu Ile Asp Glu Lys Ser Ile Ser Asn Leu Gly 35 40 45

Asp Ile His Val Ser Asp Asn Ile Leu Phe Lys Ile Val Gly Cys Asn 50 60

Asp Thr Arg Tyr Val Gly Ile Arg Tyr Phe Lys Val Phe Asp Thr Glu 65 70 75 80

Leu Gly Lys Thr Glu Ile Glu Thr Leu Tyr Ser Asp Glu Pro Asp Pro

Ser Ile Leu Lys Asp Phe Trp Gly Asn Tyr Leu Leu Tyr Asn Lys Arg 100 105 110

Tyr Tyr Leu Leu Asn Leu Leu Arg Thr Asp Lys Ser Ile Thr Gln Asn

H2

115 120 125

Ser Asn Phe Leu Asn Ile Asn Gln Gln Arg Gly Val Tyr Gln Lys Pro 130 135 140

- (2) INFORMATION FOR SEQ ID NO:4:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 143 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO:4:

As Ile Phe Ser As Thr Arg Leu Tyr Thr Gly Val Glu Val Ile Ile 1 5 10 15

Arg Lys Asn Gly Ser Thr Asp Ile Ser Asn Thr Asp Asn Phe Val Arg 20 25 30

Lys Asn Asp Leu Ala Tyr Ile Asn Val Val Asp Arg Asp Val Glu Tyr 35 40 45

Arg Leu Tyr Ala Asp Ile Ser Ile Ala Lys Pro Glu Lys Ile Ile Lys 50 55

Leu Ile Arg Thr Ser Asn Ser Asn Ser Leu Gly Gln Ile Ile Val 65 70 75 80

Met Asp Ser Ile Gly Asn Asn Cys Thr Met Asn Phe Gln Asn Asn Asn 85 90 95

Gly Gly Asn Ile Gly Leu Leu Gly Phe His Ser Asn Asn Leu Val Ala 100 105 110

Ser Ser Trp Tyr Tyr Asn Asn Ile Arg Lys Asn Thr Ser Ser Asn Gly 115 120 125

Cys Phe Trp Ser Phe Ile Ser Lys Glu His Gly Trp Gln Glu Asn 130 135 140

- (2) INFORMATION FOR SEQ ID NO:5:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1293 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: DNA (genomic)

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO:5:

TCATATACTA	ATGATAAAAT	TCTAATTTTA	TATTTTAATA	AATTATATAA	AAAATTAAA	. 60
GATAACTCTA	TTTTAGATAT	GCGATATGAA	AATAATAAAT	TTATAGATAT	CTCTGGATAT	120
GGTTCAAATA	TAAGCATTAA	TGGAGATGTA	TATATTTATT	CAACAAATAG	AAATCAATTT	180
GGAATATATA	GTAGTAAGCC	TAGTGAAGTT	AATATAGCTC	AAAATAATGA	TATTATATAC	240
AATGGTAGAT	ATCAAAATTT	TAGTATTAGT	TTCTGGGTAA	GGATTCCTAA	ATACTTCAAT	300
AAAGTGAATC	TTAATAATGA	ATATACTATA	ATAGATTGTA	TAAGGAATAA	TAATTCAGGA	360
TGGAAAATAT	CACTTAATTA	TAATAAATA	ATTTGGACTT	TACAAGATAC	TGCTGGAAAT	420
AATCAAAAAC	TAGTTTTTAA	TTATACACAA	ATGATTAGTA	TATCTGATTA	TATAAATAAA	480
TGGATTTTTG	TAACTATTAC	TAATAATAGA	TTAGGCAATT	CTAGAATTTA	CATCAATGGA	540
AATTTAATAG	ATGAAAAATC	AATTTCGAAT	TTAGGTGATA	TTCATGTTAG	TGATAATATA	600
TTATTTAAAA	TTGTTGGTTG	TAATGATACA	AGATATGTTG	GTATAAGATA	TTTTAAAGTT	660
TTTGATACGG	AATTAGGTAA	AACAGAAATT	GAGACTTTAT	ATAGTGATGA	GCCAGATCCA	720
AGTATCTTAA	AAGACTTTTG	GGGAAATTAT	TTGTTATATA	ATAAAAGATA	TTATTTATTG	780
AATTTACTAA	GAACAGATAA	GTCTATTACT	CAGAATTCAA	ACTTTCTAAA	TATTAATCAA	840
CAAAGAGGTG	TTTATCAGAA	ACCAAATATT	TTTTCCAACA	CTAGATTATA	TACAGGAGTA	900
GAAGTTATTA	TAAGAAAAA	TGGATCTACA	GATATATCTA	ATACAGATAA	TTTTGTTAGA	960
AAAAATGATC	TGGCATATAT	TAATGTAGTA	GATCGTGATG	TAGAATATCG	GCTATATGCT	1020
GATATATCAA	TTGCAAAACC	AGAGAAAATA	ATAAAATTAA	TAAGAACATC	TAATTCAAAC	1080
AATAGCTTAG	GTCAAATTAT	AGTTATGGAT	TCAATAGGAA	ATAATTGCAC	AATGAATTTT	1140
CAAAACAATA	ATGGGGGCAA	TATAGGATTA	CTAGGTTTTC	ATTCAAATAA	TTTGGTTGCT	1200
AGTAGTTGGT	ATTATAACAA	TATACGAAAA	AATACTAGCA	GTAATGGATG	CTTTTGGAGT	1260
TTTATTTCTA	AAGAGCATGG	ATGGCAAGAA	AAC			1293

(2) INFORMATION FOR SEQ ID NO:6:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1313 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: linear



(xi) SEQUENCE DESCRIPTION: SEQ ID NO:6:

GGATCCATAT GTCTTAC	ACT AACGACAAAA	TCCTGATCCT	GTACTTCAAC	AAACTGTACA	60
AAAAAATCAA AGACAAC	TCT ATCCTGGACA	TGCGTTACGA	AAACAACAAA	TTCATCGACA	120
TCTCTGGCTA TGGTTCT	AAC ATCTCTATCA	ACGGTGACGT	CTACATCTAC	TCTACTAACC	180
GCAACCAGTT CGGTATC	TAC TCTTCTAAAC	CGTCTGAAGT	AAACATCGCT	CAGAACAACG	240
ACATCATCTA CAACGGT	CGT TACCAGAACT	TCTCTATCTC	TTTCTGGGTT	CGTATCCCGA	300
AATACTTCAA CAAAGTT	AAC CTGAACAACG	AATACACTAT	CATCGACTGC	ATCCGTAACA	360
ACAACTCTGG TTGGAAA	ATC TCTCTGAACT	ACAACAAAAT	CATCTGGACT	CTGCAGGACA	420
CTGCTGGTAA CAACCAG	AAA CTGGTTTTCA	ACTACACTCA	GATGATCTCT	ATCTCTGACT	480
ACATTAATAA ATGGATC	TTC GTTACTATCA	CTAACAACCG	TCTGGGTAAC	TCTCGTATCT	540
ACATCAACGG TAACCTG	ATC GATGAAAAAT	CTATCTCTAA	CCTGGGTGAC	ATCCACGTTT	600
CTGACAACAT CCTGTTC	AAA ATCGTTGGTT	GCAACGACAC	GCGTTACGTT	GGTATCCGTT	660
ACTTCAAAGT TTTCGAC	ACT GAACTGGGTA	AAACTGAAAT	CGAAACTCTG	TACTCTGACG	720
AACCGGACCC GTCTATC	CTG AAAGACTTCT	GGGGTAACTA	CCTGCTGTAC	AACAAACGTT	780
ACTACCTGCT GAACCTG	CTC CGGACTGACA	AATCTATCAC	TCAGAACTCT	AACTTCCTGA	840
ACATCAACCA GCAGCGT	GGT GTTTATCAGA	AACCTAATAT	CTTCTCTAAC	ACTCGTCTGT	900
ACACTGGTGT TGAAGTT	ATC ATCCGTAAAA	ACGGTTCTAC	TGACATCTCT	AACACTGACA	960
ACTTCGTACG TAAAAAC	GAC CTGGCTTACA	TCAACGTTGT	TGACCGTGAC	GTTGAATACC	1020
GTCTGTACGC TGACATC	TCT ATCGCTAAAC	' CGGAAAAAAT	CATCAAACTG	ATCCGTACTT	1080
CTAACTCTAA CAACTCT	CTG GGTCAGATCA	TCGTTATGGA	CTCGATCGGT	AACAACTGCA	1140
CTATGAACTT CCAGAAC	AAC AACGGTGGTA	ACATCGGTCT	GCTGGGTTTC	CACTCTAACA	1200
ACCTGGTTGC TTCTTCT	TGG TACTACAACA	ACATCCGTAA	AAACACTTCT	TCTAACGGTT	1260
GCTTCTGGTC TTTCATC	TCT AAAGAACACG	GTTGGCAGGA	AAACTAATCT	AGA	1313

(2) INFORMATION FOR SEQ ID NO:7:

HZ

⁽i) SEQUENCE CHARACTERISTICS:(A) LENGTH: 57 base pairs

- (B) TYPE: nucleic acid (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: cDNA
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:7:

ATCGAGGGAA GGATTTCAGA ATTCGGATCC TCTAGAGTCG ACCTGCAGGC AAGCTTG

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- (2) INFORMATION FOR SEQ ID NO:8:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 5 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear

H2

- (ii) MOLECULE TYPE: peptide
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO:8:

Ile Glu Gly Arg Ile